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PATENT COOPERATION TREATY

From the INTERNATIONAL PRELIMINARY EXAMINING AUTHORITY

To:

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NOTIFICATION OF TRANSMITTAL OF

THE INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

(PCT Rule 71.1)

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Applicant's or agent's file reference

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IMPORTANT NOTIFICATION

International application No. PCT/GB2004/004390

International filing date (day/month/year)

Priority date (day/month/year)

15.10.2004

15.10.2003

Applicant

+ 44

OXFORD GENE TECHNOLOGY IP LIMITED

- The applicant is hereby notified that this International Preliminary Examining Authority transmits herewith the international preliminary report on patentability and its annexes, if any, established on the international application.
- 2. A copy of the report and its annexes, if any, is being transmitted to the International Bureau for communication to all the elected Offices.
- 3. Where required by any of the elected Offices, the International Bureau will prepare an English translation of the report (but not of any annexes) and will transmit such translation to those Offices.

4. REMINDER

The applicant must enter the national phase before each elected Office by performing certain acts (filing translations and paying national fees) within 30 months from the priority date (or later in some Offices) (Article 39(1)) (see also the reminder sent by the International Bureau with Form PCT/IB/301).

Where a translation of the international application must be furnished to an elected Office, that translation must contain a translation of any annexes to the international preliminary report on patentability. It is the applicant's responsibility to prepare and furnish such translation directly to each elected Office concerned.

For further details on the applicable time limits and requirements of the elected Offices, see Volume II of the PCT Applicant's Guide.

The applicant's attention is drawn to Article 33(5), which provides that the criteria of novelty, inventive step and industrial applicability described in Article 33(2) to (4) merely serve the purposes of international preliminary examination and that "any Contracting State may apply additional or different criteria for the purposes of deciding whether, in that State, the claimed inventions is patentable or not" (see also Article 27(5)). Such additional criteria may relate, for example, to exemptions from patentability, requirements for enabling disclosure, clarity and support for the claims.

Name and mailing address of the international preliminary examining authority:

European Patent Office - P.B. 5818 Patentlaan 2 NL-2280 HV Rijswijk - Pays Bas Tel. +31 70 340 - 2040 Tx: 31 651 epo nl Fax: +31 70 340 - 3016

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PATENT COOPERATION TREATY

PCT

INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

(Chapter II of the Patent Cooperation Treaty)

(PCT Article 36 and Rule 70)

Applicant's or agent's file	reference					
P035667WO:CJM FOR FURTHER A		CTION See Form PCT/IPEA/416				
International application PCT/GB2004/00439		International filing date 15.10.2004	(day/month/year)	Priority date (day/month/year) 15.10.2003		
International Patent Clas B01J19/00, H01L51		tional classification and	PC			
Applicant OXFORD GENE TECHNOLOGY IP LIMITED						
This report is the international preliminary examination report, established by this International Preliminary Examining Authority under Article 35 and transmitted to the applicant according to Article 36.						
2. This REPORT of	2. This REPORT consists of a total of 6 sheets, including this cover sheet.					
3. This report is als						
a. 🛭 sent to th						
{ and/c	ts of the description or sheets containing inistrative Instruction	g rectifications authori	ngs which have beer ized by this Authority	n amended and are the basis of this report (see Rule 70.16 and Section 607 of the		
beyo	ts which supersed nd the disclosure i lemental Box.	e earlier sheets, but w n the international app	hich this Authority co dication as filed, as in	nsiders contain an amendment that goes adicated in item 4 of Box No. I and the		
sequence	e listing and/or table	ureau only) a total of (i es related thereto, in o Listing (see Section 80	computer readable for	nber of electronic carrier(s)) , containing a rm only, as indicated in the Supplemental re Instructions).		
4. This report conta	nins indications rela	ating to the following it	tems:			
☑ Box No. I	Basis of the opini	ion				
☐ Box No. II	Priority					
☐ Box No. III	Non-establishme	nt of opinion with rega	ard to novelty, invention	e step and industrial applicability		
☐ Box No. IV	Lack of unity of ir	nvention				
⊠ Box No. V	Reasoned statem applicability; citat	nent under Article 35(2 tions and explanations	2) with regard to nove supporting such stat	Ity, inventive step or industrial ement		
☐ Box No. VI	Certain documen					
☐ Box No. VII	Certain defects in	n the international app	lication			
⊠ Box No. VIII	Certain observati	ions on the internation	al application			
Date of submission of the demand		Date of completion of	this report			
13.06.2005			31.01.2006			
Name and mailing address of the international preliminary examining authority:			Authorized Officer			
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INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

International application No. PCT/GB2004/004390

_	Box No. I Basis of the repor	1	
1		nis report is based on the international application in the language in which it was	
	international search (un □ publication of the internation	nslations from the original language into the following language, translation furnished for the purposes of: der Rules 12.3 and 23.1(b)) ational application (under Rule 12.4) r examination (under Rules 55.2 and/or 55.3)	
2. With regard to the elements* of the international application, this report is based on (replacement have been furnished to the receiving Office in response to an invitation under Article 14 are referred report as "originally filed" and are not annexed to this report):			
	Description, Pages		
	1-23	as originally filed	
	Claims, Numbers		
	1-10	as originally filed	
	11-41	filed with telefax on 30.01.2006	
	Drawings, Sheets		
	1/5-5/5	as originally filed	
	☐ a sequence listing and/or ar	ny related table(s) - see Supplemental Box Relating to Sequence Listing	
3.	☐ The amendments have rest ☐ the description, pages ☐ the claims, Nos. ☐ the drawings, sheets/figs ☐ the sequence listing (specified any table(s) related to see	; ecify):	
1.	☐ This report has been estable had not been made, since they he supplemental Box (Rule 70.2(c)) ☐ the description, pages ☐ the claims, Nos. ☐ the drawings, sheets/figs ☐ the sequence listing (special any table(s) related to see	ecify):	
	* If item 4 applies, so	ome or all of these sheets may be marked "superceded "	

INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

International application No. PCT/GB2004/004390

Box No. V Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N)

Yes: Claims

1-41

No: Claims

Inventive step (IS)

Yes: Claims

1-41

No: Claims

Industrial applicability (IA)

Yes: Claims

1-41

No: Claims

2. Citations and explanations (Rule 70.7):

see separate sheet

Box No. VIII Certain observations on the international application

The following observations on the clarity of the claims, description, and drawings or on the question whether the claims are fully supported by the description, are made:

see separate sheet

Reference is made to the following documents:

- D1: WO 03/020415 A2 (ISIS INNOVATION LIMITED; SOUTHERN, EDWIN, MELLOR; EGELAND, RYAN, D) 13 March 2003 (2003-03-13)
- D2: US-B1-6 630 359 (CAILLAT PATRICE ET AL) 7 October 2003 (2003-10-07)
- D3: US-A-6 093 302 (MONTGOMERY ET AL) 25 July 2000 (2000-07-25)
- D4: WO 01/43870 A (MOTOROLA INC; SHI, SONG; ZHANG, PEIMING; MARACAS, GEORGE; MARACAS, GEO) 21 June 2001 (2001-06-21)
- D5: WO 98/01758 A (NANOGEN, INC) 15 January 1998 (1998-01-15)

Re Item V

Subject to the remark about independent claim 13 (which is unclear) in Item VIII, the following is observed.

1. Document D1 discloses a method and a device for electrochemically treating a substrate with electrodes producing first and second quenchable redox products. The electrodes are individually addressable (see step (b) in claim 20) and there can be one common counter electrode (see claim 25). The subject-matter of independent claims 1 and 13 differs from D1 in having a common first electrode arranged to define cells therein.

Document D2 (see passages cited in the Search Report) describes a system and method for electrochemically treating a substrate. It has an arrangement of a common first electrode ((9b) or (29) in Figures 1 and 2) arranged to define cells therein and individually addressable second electrodes. In column 1, lines 26-38, it becomes clear that this document is concerned with the demand for systems enabling the chemical or biological analysis with a very large number of points (i.e., high resolution). D2, however changes the surface of the addressable electrodes itself and not of a substrate above it. The subject-matter of claims 1 and 13 differs in that the electrodes and the substrate are separated by the electrolyte. It is not suitable for patterning several substrates

Document D3 discloses a method and a device for electrochemically treating a substrate with addressable electrodes and "getter" electrodes arranged to define cells around the addressable electrodes (see for example Figure 36). The electrolyte (usually

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water) is such that the second redox product (H⁺) is quenchable by the first redox product (OH⁻). The "getter" electrode may be used in conjunction with a scavenging solution and has the function of scavenging electrochemically generated reagents that may diffuse away from the electrode. The subject-matter of claim 1 differs from D3 in that the substrate appears to be separated from the device (the device faces the substrate), the first electrode is "common", and that the getter electrodes appears to be not necessarily the counter electrodes.

The subject-matter of independent claims 1 and 13 is therefore <u>novel</u> (Article 33(2) PCT).

Document D1 is at present considered to be the closest prior art.

The difference with D1 seems to be twofold: the electrode is "common" and it defines cells. This seems to lead to a simple arrangement allowing modification of a substrate with improved resolution.

The problem to be solved appears to be the provision of a means for repeatedly patterning different substrates (excluding D2 and D3 which deal with electrodes on a substrate as closest prior art) comprising a simple arrangement allowing improved resolution.

It is not straightforward to at the same time provide improved resolution and a simple device. The arrangement of claims 1 and 13 surprisingly achieves this goal.

The subject-matter of claims thus 1 and 13 appears to involve an inventive step (Article 33(3) PCT).

The dependent claims as such then also meet the requirements of the PCT with respect to novelty and inventive step.

Although the category and dependency of claims 31-38 is unclear, they would also be novel and inventive if they would properly depend on claim 1.

Independent claims 39 and 41, as well as dependent claim 40, appear to be novel and

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International application No.

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inventive for the same reasons as mentioned for claim 1.

Re item VIII

1. Independent claim 13 is unclear (Art. 6 PCT). It defines the device, *inter alia*, by means of the electrolyte which is in contact with the substrate <u>during use</u>. This throws doubt upon the category of the claim (use-claim or entity claim?) and (therefore) the scope of desired protection.

If "during use" had not been in the claim, it would have been acceptable under Art. 6 PCT. Basis for the omission of "during use" appears to be present on page 2, lines 23-25.

2. Claims 21,29-41 do not comply with the requirements of Article 6 PCT.

Claims 21,29 and 30 are formulated in such broad terms, referencing to (parts of) other unspecified devices that the scope of protection could not be clearly determined.

For claims 31-38 both the category and the dependency, and even if they are to be considered as independent or dependent, are not well defined.

The combination of independent claims 1, 39 and 41 leads to a multiplicity of independent claims in the same category. Although these claims have been drafted as separate independent claims, they appear to relate effectively to the same subject-matter and to differ from each other only with regard to the definition of the subject-matter for which protection is sought and/or in respect of the terminology used for the features of that subject-matter. The aforementioned claims therefore lack conciseness and as such do not meet the requirements of Article 6 PCT.

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- 11. The method of any preceding claim, wherein the substrate material is impermeable.
- 12. The method of any preceding claim, wherein the substrate material is a glass, a plastic, a metal, a semiconductor, a silicon oxide or a gel.
- 13. A device for electrochemically modifying a substrate, the device having:
 - (i) a common first electrode arranged to define cells therein; and
 - (ii) a phrality of individually addressable second electrodes,

wherein:

- (a) a phurality of the cells contain individually addressable second electrodes;
- (b) the common first electrode and the plurality of second electrodes are in contact with an electrolyte;
- (c) the electrolyte is in contact with the substrate during use, the substrate being the surface of a material which is separate from and facing the electrodes; and
- (d) the electrolyte is such that the common first electrode is able to generate a first redox product, the second electrode is able to generate a second redox product, and the second redox product is quenchable by the first redox product.
- 14. The device of claim 13, wherein the common first electrode is a cathode and the second electrodes are anodes.
- 15. The device of any one of claims 13-14, wherein the common first electrode is singly addressable such that the potential of the first electrode may be altered by addressing a single connection.
- 20 16. The device of any one of claims 13-14, wherein the common first electrode is a bus line which is neither switchable nor addressable.
 - 17. The device of any one of claims 13-16, wherein the common first electrode has a geometry in which there is a substantially regular pattern, such as a grid, a net, a honeycomb, a series of intersecting circles of other tessellating shapes.
- 18. The device of any one of claims 13-17, wherein the arrangement of the common first electrode results in at least $n \times 10^6$ cells, where n is 0.5 or more.
 - 19. The device of any one of claims 13-18, wherein the arrangement of the common first electrode is such that the distance between the centre of a cell and the centre of at least one of its immediate neighbour cells is less than 0.5 mm.
- 20. The device of any one of claims 13-19, wherein the second electrodes are addressed using a direct connection from an electrode to a bond pad on the perimeter of the device, by CMOS switching circuitry, or by transistor-based circuitry.

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- 21. The device of claim 20, wherein the second electrodes are addressed using TFT circuitry.
- 22. The device of any one of claims 13-21, wherein the distance between the centre of a second electrode and the centre of at least one of its neighbouring second electrodes is less than 0.5 mm.
- 23. The device of claim any one of claims 13-22, comprising at least $n \times 10^6$ second electrodes, where n is 0.5 or more.
- 24. The device of any one of claims 13-23, wherein the ratio of the number of cells to the number of second electrodes is less than 1.
- 25. The device of any one of claims 13-23, wherein the ratio of the number of cells to the number of second electrodes is more than 1.
- 26. The device of any one of claims 13-23, wherein the ratio of the number of cells to the number of 10 second electrodes is substantially 1.
 - 27. The device of claim any one of claims 13-26, wherein the common first electrode and second electrodes are made from materials independently selected from indium tin oxide, iridium, platinum, palladium, gold, silver, copper, nickel, zinc, titanium, tungsten, aluminium and alloys of these metals.
 - 28. The device of claim 27, wherein the common first electrode and second electrodes comprise a coating of iridium on another material.
 - 29. The device of any one of claims 13-28, wherein the support on which the second electrodes are positioned and the circuitry connecting the second electrodes are as found in liquid crystal display devices.
 - 30. The device of any one of claims 13-29, wherein incorporated into a flow cell arrangement.
 - 31. The method of any one of claims 1 to 12, or the device of any one of claims 13 to 30, for use in deprotecting a substrate in specific patterns.
 - 32. The method or device of claim 31, for use in removing protecting groups from specific regions of a substrate to leave a pattern of deprotected groups, such that subsequent exposure of the 25 deprotected groups to a reactant allows the deprotected groups to react with the reactant.
 - 33. The method or device of claim 31, for use in combinatorial synthesis.
 - 34. The method or device of claim 31, for use in the synthesis of libraries of small organic compounds bound to a surface.
 - 35. The method or device of claim 31, for use in synthesising polymers.

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- 36. The method or device of claim 34, for use in synthesising polynucleotides, polysaccharides, or polypeptides.
- 37. The method or device of claim 31, for etching of the substrate.
- 38. The method or device of claim 31, for use in the production of organic LED materials.
- 5. 39. A method for preparing an array of polynucleotides comprises the steps of:
 - (1) providing a substrate having protecting groups on its surface and which faces the device of any one of claims 13 to 30 and is in contact with the electrolyte;
 - (2) switching a first set of second electrodes so as to generate a second redox product which exposes deprotected groups by removing a set of protecting groups from the substrate;
 - (3) coupling a nucleotide to the set of deprotected groups, the nucelotide comprising a protecting group; and
 - (4) repeating the sequence of steps (2) and (3) until the desired array has been generated.
 - 40. The method of claim 39, wherein the second redox product is a proton and the protecting group is an acid-labile protecting group which protects a furanyl hydroxyl group.
- 41. A method for guiding a reagent along the face of a substrate in a desired path, wherein a device of any one of claims 13 to 30 is used to create redox products along the desired path, and wherein the reagent moves into region(s) where the redox products are created.

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